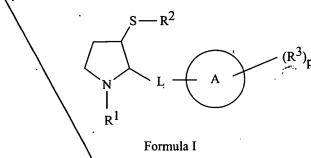
IN THE CLAIMS:

Please cancel claims 2, 4, 5, and 6 from the present application without disclaimer or prejudice.

Please amend claims 1, 3, 7, 8, 9, 11, and 12 as follows:

Claim 1:

1. A compound of the Formula I



wherein:

R¹ is selected from H; -C₁₋₄alkyl; -CO-C₁₋₄alkyl; -CO-O-C₁₋₄alkyl;

-CO-O-C₂₋₄alkenyl; -C₁₋₄alkylene-CONR⁴R⁵ (wherein R⁴ and R⁵ are independently selected from H and C₁₋₄alkyl); -C₁₋₄alkylene-COOR⁶ (wherein R⁶ is selected from H and C₁₋₄alkyl); -C₁₋₃alkylene-Ph and -CO-O(CH $_2$)_nPh wherein the phenyl groups in -C₁₋₃alkylene-Ph and -CO-O(CH $_2$)_nPh are optionally substituted by R^a and/or R^b and R^a and R^b are independently selected from C₁₋₄alkyl, halogen, hydroxy, C₁₋₄alkoxy, C₁₋₄alkanoyl, C₁₋₄alkanoyloxy, amino, C₁₋₄alkylamino, di(C₁₋₄alkyl)amino, C₁₋₄alkanoylamino, nitro, cyano, carboxy, carbamoyl, C₁₋₄alkoxycarbonyl, thiol, C₁₋₄alkylsulfanyl, C₁₋₄alkylsulfinyl,C₁₋₄alkylsulfonyl and sulfonamido; and n=0-4;

R² is selected from H; -C₁₋₄alkyl; -COC₁₋₄alkyl; and -COOC₁₋₄alkyl; and -C₁₋₃alkylene-Ph optionally substituted on the phenyl ring by R^a and/or R^b;

R³ is selected from H; OH; CN; CF3; NO2; -C₁₋₄ alkyl; -C₁ alkylene-R⁷;

-C₂₋₄alkenylene-R⁷; -C₂₋₄alkynylene-R⁷; R⁷; OR⁷ (where R⁷ is selected from phenyl, naphthyl, a 5-10 membered monocyclic or bicyclic heteroaryl ring containing upto 5

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heteroatoms selected from O,N and S and any aryl ring in R^7 is optionally substituted by R^a and/or R^b), C_{2-4} alkenyl; halogen; -(CH_2) $_n$ COOR 8 (where n=0-3 and R^8 represents H, C_{1-4} alkyl, or C_{2-4} alkenyl); -CONR 9 R 10 (where R^9 and R^{10} independently represent H, C_{1-4} alkyl, C_{2-4} alkenyl, -O- C_{1-4} alkyl, -O- C_{2-4} alkenyl or -C1-3alkylenePh (wherein Ph is optionally substituted by R^a and R^b as hereinabove defined);-CON(R^{11})OR 12 (where R^{11} and R^{12} independently represent H, C_{1-4} alkyl or C_{2-4} alkenyl);

-CONR¹³-CR^{13a} R¹⁴-COOR¹⁷, where R¹³ and R^{13a} are independently H or C₁₋₄alkyl, R¹⁷ is H or C₁₋₆alkyl, R¹⁴ is selected from the side chain of a lipophilic amino acid, carbamoylC₁₋₄alkyl, N-(monoC₁₋₄alkyl)carbamoylC₁₋₄alkyl and N-(diC₁₋₄alkyl)carbamoylC₁₋₄alkyl) having L or D configuration at the chiral alpha carbon in the corresponding free amino acid; a lactone of formula:

-con O

C₁₋₄alkyl monosubstituted on carbon with =N-OH;

a group of Formula -X-R¹⁵ (where X is selected from 0, CO, CH₂, S, SO, SO₂ and R¹⁵ is selected from C₁₋₆alkyl, phenyl, naphthyl, a 5-10 membered monocyclic or bicyclic heteroaryl ring containing upto 5 heteroatoms selected from O,N and S and any aryl ring in R¹⁵ is optionally substituted by R^a and/or R^b;

p is 0-3 in which R³ values can be the same or different;

L is a linking moiety selected from the following groups written from left to right in Formula I:

Br

(wherein the piperazine and perhydro-1,4-diazepine rings are optionally substituted);

-CO-NR¹⁶-; -CH₂-NR¹⁶-; -CH₂S-; -CH₂O-; -CH₂-CHR¹⁶; -CH=CR¹⁶-; -CH₂NR¹⁶-T-;

 $-CH_{2}NR^{16}-SO_{2}-; -CH_{2}-NR^{16}-CO-T^{1}-; -CO-NR^{16}-T-; -CH_{2}S-T-; -CH_{2}O-T- \ (where \ R^{16} \ is \ R^{16}-R^$

selected from H, C₁₋₄alkyl, C₁₋₄alkylene-Z, -CO-C₁₋₄alkylene-Z, -CO-C₁₋₆alkyl, -COZ,

Z and Z is selected from -O C₁₋₄alkyl, phenyl, naphthyl, a 5-10 membered monocyclic or bicyclic heteroaryl ring containing upto 5 heteroatoms selected from O, N and S and

anv arvl ring in R¹⁶ is optionally substituted by R^a and/or R^b as hereinabove defined;

where, T represents -(CH₂)m- where n is 1-4 and T is optionally monosubstituted with

any value of R¹⁶ other than H; and

where T¹ represents -(CH₂)m¹- wherein m¹ is 0-4 and T is optionally monosubstituted

with any value of R¹⁶ other than H);

A is selected from phenyl; naphthyl; a 5-10 membered monocyclic or bicyclic heteroaryl ring containing upto 5 heteroatoms where the heteroatoms are independently selected from O, N & S;

or a -S-S- dimer thereof when R²=H; or a N-oxide thereof; or a pharmaceutically acceptable salt, prodrug or solvate thereof.

Claim 3:

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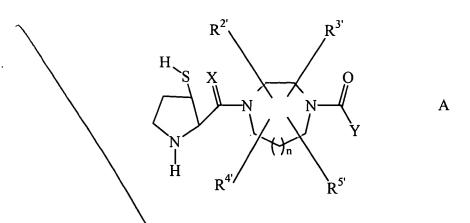
3. A compound according to claim 1 wherein A is phenyl or naphthyl.

Claim 7:

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7. A compound of the formula A:

By Cont



wherein:

X is O or H₂;

n is 0 or 1;

t is 1 to 4;

R^{2'}, R^{3'}, R^{4'}, and R^{5'} are independently selected from: H; C₁₋₈alkyl, alkenyl, alkynyl, aryl, heterocycle, -CO-NR^{6'}R^{7'} or -CO-OR^{6'}, unsubstituted or substituted with one or more of:

- 1) aryl or heterocycle, unsubstituted or substituted with:
 - a. C₁₋₄alkyl,
 - b. (CH₂)tOR⁶,
 - c. $(CH_2)_t NR^{6'}R^{7'}$,
 - d. halogen,
- 2) C₃₋₆cycloalkyl,
- 3) $OR^{6'}$,
- 4) $SR^{6'}$, $S(O)R^{6'}$, $SO_2R^{6'}$,
- 5) $-NR^{6'}R^{7'}$,
- 6) $-NR^{6'}-CO-R^{7'}$,
- 7) $-NR^{6'}-CO-NR^{7'}R^{8'}$,

B4 Cont

- 8) $-0-CO-NR^{6'}R^{7'}$,
- 9) \-O-CO-OR^{6'},
- 10) $\frac{1}{2} NR^{6'}R^{7'}$,
- 11) $-SO_{2}NR^{6'}R^{7'}$,
- 12) -NR⁶'-\$O₂-R⁷',
- 13) $-CO-R^{6'}$, α
- 14) -CO-OR⁶;

and any two of R², R³, R⁴, and R⁵ are optionally attached to the same carbon atom; Y is aryl, heterocycle, unsubstituted or substituted with one or more of:

- 1) C₁₋₄alkyl, unsubstituted or substituted with:
 - a. C₁₋₄alkoxy,
 - b. $NR^{6'}R^{7'}$,
 - c. C₃₋₆cycloalkyl,
 - d. aryl or heterocycle,
 - e. HO,
- 2) aryl or heterocycle,
- 3) halogen,
- 4) $OR^{6'}$,
- 5) $NR^{6'}R^{7'}$,
- 6) CN
- 7) NO₂, or
- 8) CF₃;

R⁶, R⁷ and R⁸ are independently selected from: H; C₁-talkyl, C₃-6cycloalkyl, heterocycle, aryl, aroyl, heteroaroyl, arylsulfonyl, heteroarylsulfonyl, unsubstituted or substituted with:

a) C₁₋₄alkoxy,

By Cont

- b) aryl or heterocycle,
- c) hà(ogen,
- d) HO,
- e) -CO-R^S
- f) -SO₂R^{9'}, wherein

R^{6'} and R^{7'} may be joined in a ring, and

R⁷ and R⁸ may be joined in a ring;

R⁹ is C₁₋₄alkyl or aralkyl;

a pharmaceutically acceptable salt thereof.

Claim 8:

- 8. A compound according to claim 1 which is any one of the following individual compounds or a pharmaceutically acceptable salt thereof:
- (2S)-2-{2-benzyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-{2-benzyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-({2-phenyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-({2-phenyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid;
- (2S)-2-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl)amino]-naphthalenè-1-carbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-({3-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl)amino]-naphthalene-1-carbonyl}-amino)-4-methylsulfanylbutyric acid;
- (2S)-2-({-3-phenyl-5[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;

By Cont

(2S)-2-(\darkappa-3-phenyl-5[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid;

(cis)-2-[{N-(4-nethoxybenzyl)- N-(naphthalen-1-ylmethylamino}-methyl]-pyrrolidine-3-thiol;

N-(naphthalen-1-ylmethyl)-N-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl)-pentanamide; N-(naphthalen-1-ylmethyl)-N-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl)-2-(pyridin-3-yl)-acetamide :

N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-3-methyl-N-(2-naphthalen-1-ylethyl)butyramide;

N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-N-(2-naphthalen-1-yl-ethyl)-2-pyridin-3-yl-acetamide :

(cis)-2-{[(3-methoxypropyl)-(2-naphthalen-1-ylethyl)amino]methyl}- pyrrolidine-3-thiol; N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-2-(4-methoxy-phenyl)-N-(2-naphthalen-2-yl-

ethyl)-acetamide;

(cis)-2-{[(2-(4-methoxyphenyl)ethyl)-(2-naphthalen-1-ylethyl)amino] methyl}- pyrrolidine-3-thiol;

 $N-(2,2-diphenyl-ethyl)-N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-3-methyl-butyramide\ ;\\ N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-3,3-dimethyl-N-(2-naphthalen-2-yl-ethyl-ethyl-N-(2-naphthalen-2-yl-ethyl-ethyl-naphthalen-2-yl-ethyl-N-(2-naphthalen-2-yl-ethyl-naphthalen$

butyramide;

N-(2,2-diphenyl-ethyl)-N-[(cis)-3-sulfanyl-pyrrolidin-2\sylmethyl)-3,3-dimethyl-butyramide;

(2S)-2-{3-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-(3-methoxy-propyl)-amino]-benzoylamino}-4-methylsulfanyl-butyric acid;

N-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-3,3-dimethyl-N-(2-naphthalen-1-yl-ethyl)-butyramide;

- (2S)-4-carbamoyl-2-({2-phenyl-5-[(cis)-3-sulfanyl-pyrrolidin-2-x|methyl)-amino]-phenylcarbonyl}-amino)-butyric acid;
- (2S)-4-carbamoyl-2-({2-phenyl-5-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-butyric acid methyl ester;
- 2-(3-pyridyl)-N-(2,2-diphenyl-ethyl)-N-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl)- acetamide; 6-methoxy-1-oxido-N-(2,2-diphenyl-ethyl)-N-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl]-pyridine-3-carboxamide;

By Cont

- N-(naphthyl-1-yl-ethyl)-N-[(cis)-3-sulfanylpyrrolidin-2yl-methyl)-thiazole-5-carboxamide; 6-methoxy-1-oxido-N-(naphthyl-1-yl-ethyl)-N-[cis)-3-sulfanylpyrrolidin-2-ylmethyl]-pyridine-3-carboxamide;
- (2S)-2-{2-benzyl-4-[(cis)-3-sulfanyl-pyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanyl-butyric acid;
- (2S)-2-{2-benzyl-5-\((cis)-3-sulfanylpyrrolidin-2-ylmethyl)amino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-benzyl-4-[(cis)-8-sulfanylpyrrolidin-2-ylmethyl)amino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-phenethyl-5-[(trans)-3-sulfanylpyrrolidin-2-ylmethylaminobenzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{phenethyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-benzyl-5-[(trans)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-(phenethyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-(4-methylphenylethynyl)-4-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid;
- (2S)-2-{2-benzyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid isopropyl ester;
- (2S)-2-{2-benzyl-4-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-{2-benzyl-4-[(trans)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-{2-benzyl-5-[(trans)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-{2-phenyl-5-[(trans)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;
- (2S)-2-{2-phenyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}\d-methylsulfanylbutyric acid methyl ester;

By Casa cont

(2S)-2-(2-benzyl-5-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-{2-(4-methylphenethyl)-4-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino]-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-{2-(4-methylphenylethynyl)-4-[(cis)-3-sulfanylpyrrolidin-2-ylmethylamino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-(2-methoxyethyl)-1-[(cis)-3-sulfanylpyrrolidin-2-ylmethyl]-4-(naphth-1-oyl)piperazine;

(cis)-2-[N-isovaleryl-N-(2-(napth-1-yl)ethyl)aminiomethyl]-3-sulfanylpyrrolidine;

(cis)-2-[N-(3-pyridylacetyl)-N-(\(\pi\)aphth-1-yl)ethyl)aminomethyl]-3-sulfanylpyrrolidine;

(cis)-2-[N-1-oxido-6-methoxypyridin-3-ylcarbonyl) -N-(naphth-1-yl)ethyl)aminomethyl]-3-sulfanylpyrrolidine;

(cis)-2-[N-thiazol-5-ylcarbonyl) -N-(naphth-1-yl)ethyl)aminomethyl]-3-sulfanylpyrrolidine;

(2S)-2-[2-(4-fluorophenethyl)-4-[(cis)-3-sulfanyl)-pyrrolidin-2-

ylmethylamino)benzoylamino]-4-methylsulfanylbutyric acid;

methyl (2S)-2-[2-(4-fluorophenethyl)-4-[(cis)-3-sulfanylpyrrolidin-2-

ylmethylamino)benzoylamino]-4-methylsulfanylbutyrate;

(2S)-2-[2-(4-fluorophenethyl)-4-((2R,3R)-3-sulfanyl-pyrrolidin-2-

ylmethylamino)benzoylamino]-5-methylsulfanylbutyric acid;

(2S)-2-{2-Benzyl-5-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-benzoylamino}-4-methylsulfanylbutyric acid methyl ester ;

(2S)-2-{2-Benzyl-5-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-benzoylamino}-4-methylsulfanylbutyric acid ;

(2S)-2-({2-phenyl-5-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-({2-phenyl-5-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid;

(2S)-2-({3-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-naphthalene-1-carbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-({3-[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-naphthalene-1-carbonyl}-amino)-4-methylsulfanylbutyric acid ;

Cont Cont

(2S)-2-((-3-phenyl-5[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid methyl ester;

(2S)-2-({-3-phenyl-5[([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-4-methylsulfanylbutyric acid;

(2R,3R)-2-[{N-(4-methoxybenzyl)- N-(naphthalen-1-ylmethyl)-amino}-methyl]-pyrrolidine-3-thiol;

N-(naphthalen-1-ylmethyl)-N-([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-pentanamide;

N-(naphthalen-1-ylmethyl)-N-([2R,3R]-3-sulfanylpyrrolidin-2-ylmethyl)-2-(pyridin-3-yl)-acetamide :

N-((2R,3R)-3-sulfanyl-pyrrolidin-2-ylmethyl)-3-methyl-N-(2-naphthalen-1-ylethyl)butyramide;

N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-N-(2-naphthalen-1-yl-ethyl)-2-pyridin-3-yl-acetamide :

(2R,3R)-2-{[(3-Methoxypropyl)-(2-naphthalen-1-ylethyl)amino]methyl}- pyrrolidine-3-thiol;

N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-2-(4-methoxy-phenyl)-N-(2-naphthalen-2-ylethyl)-acetamide;

(2R,3R)-2-{[(2-(4-Methoxyphenyl)ethyl)-(2-naphthalen-1-ylethyl)amino] methyl}-pyrrolidine-3-thiol;

N-(2,2-Diphenyl-ethyl)-N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-3-methyl-butyramide

N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-3,3-dimethyl-N-(2-naphthalen-2-yl-ethyl)-butyramide;

N-(2,2-Diphenyl-ethyl)-N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-3,3-dimethyl-butyramide;

(2S)-2-{3-[([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-(3-methoxy-propyl)-amino]-benzoylamino}-4-methylsulfanyl-butyric acid;

N-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-3,3-dimethyl-N-(2-naphthalen-1-yl-ethyl)-butyramide;

(2S)-4-carbamoyl-2-({2-phenyl-5-[([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-butyric acid;

By Cont

(2S)-4-carbamoyl-2-({2-phenyl-5-[([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-amino]-phenylcarbonyl}-amino)-butyric acid methyl ester;

2-(3-pyridyl)-N-(2,2-diphenyl-ethyl)-N-((2R,3R)-3-sulfanylpyrrolidin-2-ylmethyl)-acetamide;

6-methoxy-1-oxido-N-(2,2-diphenyl-ethyl)-N-((2R,3R)-3-sulfanylpyrrolidin-2-ylmethyl)-pyridine-3-carboxamide;

N-(naphthyl-1-yl-ethyl)-N-([2R,3R]-3-sulfanylpyrrolidin-2yl-methyl)-thiazole-5-carboxamide;

6-methoxy-1-oxido-N-(naphthyl-1-yl-ethyl)-N-((2R,3R)-3-sulfanylpyrrolidin-2-ylmethyl)-pyridine-3-carboxamide;

(2S)-2-{2-benzyl-4-[([2R,3R]3-sulfanyl-pyrrolidin-2-ylmethyl)-amino]-benzoylamino}-4-methylsulfanyl-butyric acid; and

(2S)-2-(2-methoxy-ethyl)-1-([2R,3R]-3-sulfanyl-pyrrolidin-2-ylmethyl)-4-naphthoyl-piperazine.

Claim 9:

9. A pharmaceutical composition which comprises a compound according to any one of claims 1, 3, 7, or 8 and a pharmaceutically-acceptable carrier.

Claim 11:



11. A compound according to any one of claims 1, 3, 7 or 8 for use as a medicament.

Claim 12:

12. A compound according to any one of claims 1, 3, 7 or 8 for use in the preparation of a medicament for treatment of a disease mediated through farnesylation of mutant ras.

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Please add claims 14-17 as follows: